

chain nodes :

7 14 17

ring nodes :

1 2 3 4 5 6 8 9 10 11 12 13 19 20 21 22 23 24

chain bonds :

2-7 5-9 6-21 7-14

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 8-9 8-13 9-10 10-11 11-12 12-13
19-20 19-24 20-21 21-22 22-23 23-24

exact/norm bonds :

1-2 1-6 2-3 2-7 3-4 4-5 5-6 5-9

exact bonds :

6-21 7-14

normalized bonds :

8-9 8-13 9-10 10-11 11-12 12-13 19-20 19-24 20-21 21-22 22-23
23-24

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:CLASS 8:Atom 9:Atom
10:Atom 11:Atom 12:Atom 13:Atom 14:CLASS 17:CLASS 18:CLASS
19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 24:Atom

10/073960

=> s l1

SAMPLE SEARCH INITIATED 18:45:59 FILE 'REGISTRY'
SAMPLE SCREEN SEARCH COMPLETED - 76 TO ITERATE

100.0% PROCESSED 76 ITERATIONS
SEARCH TIME: 00.00.01

0 ANSWERS

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**
PROJECTED ITERATIONS: 997 TO 2043
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 18:46:12 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 2053 TO ITERATE

100.0% PROCESSED 2053 ITERATIONS
SEARCH TIME: 00.00.01

4 ANSWERS

L3 4 SEA SSS FUL L1

=> d l3 1-4

10/073960

L3 ANSWER 1 OF 4 REGISTRY COPYRIGHT 2002 ACS

RN 457601-25-1 REGISTRY

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)-, dihydrate
(9CI) (CA INDEX NAME)

OTHER NAMES:

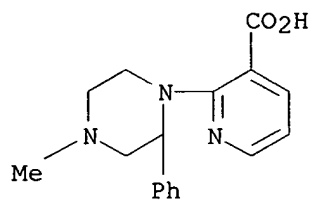
CN 1-(3-Carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine dihydrate

MF C17 H19 N3 O2 . 2 H2 O

SR CA

LC STN Files: CA, CAPLUS, CASREACT, USPATFULL

CRN (61338-13-4)



● 2 H₂O

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

10/073960

L3 ANSWER 2 OF 4 REGISTRY COPYRIGHT 2002 ACS

RN 343626-55-1 REGISTRY

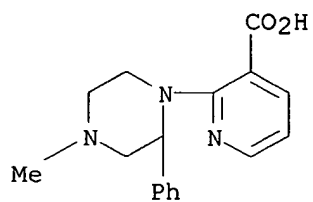
CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)-, potassium
salt (9CI) (CA INDEX NAME)

MF C17 H19 N3 O2 . K

SR CA

LC STN Files: CA, CAPLUS, USPAT2, USPATFULL

CRN (61338-13-4)



● K

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

10/073960

L3 ANSWER 3 OF 4 REGISTRY COPYRIGHT 2002 ACS

RN 125967-25-1 REGISTRY

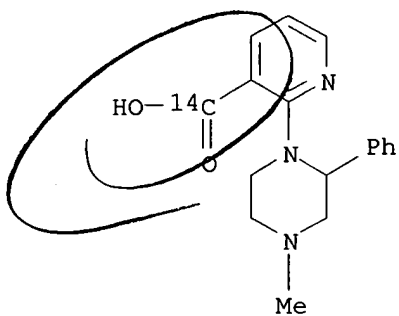
CN 3-Pyridinecarboxylic-14C acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI)
(CA INDEX NAME)

FS 3D CONCORD

MF C17 H19 N3 O2

SR CA

LC STN Files: CA, CAPLUS, CASREACT



1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

10/073960

L3 ANSWER 4 OF 4 REGISTRY COPYRIGHT 2002 ACS

RN 61338-13-4 REGISTRY

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA INDEX NAME)

OTHER NAMES:

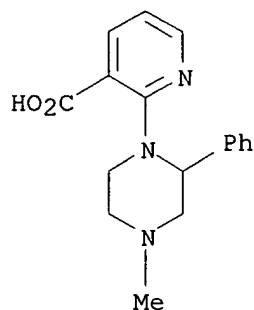
CN 1-(3-Carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine

FS 3D CONCORD

MF C17 H19 N3 O2

CI COM

LC STN Files: BEILSTEIN*, CA, CAPLUS, CASREACT, IFICDB, IFIPAT, IFIUDB, USPAT2, USPATFULL
(*File contains numerically searchable property data)



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

6 REFERENCES IN FILE CA (1962 TO DATE)

6 REFERENCES IN FILE CAPLUS (1962 TO DATE)

10/073960

=> s l3 and ?hydrate?/csn

'CSN' IS NOT A VALID FIELD CODE

LEFT TRUNCATION IGNORED FOR '?HYDRATE?' FOR FILE 'REGISTRY'

0 HYDRATE?/CSN

L4 0 L3 AND ?HYDRATE?/CSN

Left truncation is not valid in the specified search field in the specified file. The term has been searched without left truncation.

Examples: '?TERPEN?' would be searched as 'TERPEN?' and '?FLAVONOID' would be searched as 'FLAVONOID.'

If you are searching in a field that uses implied proximity, and you used a truncation symbol after a punctuation mark, the system may interpret the truncation symbol as being at the beginning of a term. Implied proximity is used in search fields indexed as single words, for example, the Basic Index.

=> help field codes

HELP FOR 'FIELD CODES' IS NOT AVAILABLE

For information about help messages available in all files, enter "HELP MESSAGES". For information about help messages available for the current file, enter "HELP DIRECTORY". For a list of commands, enter "HELP COMMANDS".

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

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148.33

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FILE COVERS 1907 - 17 Dec 2002 VOL 137 ISS 25

FILE LAST UPDATED: 16 Dec 2002 (20021216/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

CAS roles have been modified effective December 16, 2001. Please check your SDI profiles to see if they need to be revised. For information on CAS roles, enter HELP ROLES at an arrow prompt or use the CAS Roles thesaurus (/RL field) in this file.

=> s l3

L5 7 L3

10/073960

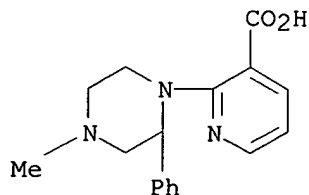
=> d 15 1-7 bib abs hitstr

10/073960

L5 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2002 ACS
AN 2002:695977 CAPLUS
DN 137:216962
TI Methods for the preparation of mirtazapine intermediates
IN Metzger, Leonid; Wize, Shlomit
PA Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals USA, Inc.
SO PCT Int. Appl., 12 pp.
CODEN: PIXXD2
DT Patent
LA English
FAN.CNT 1

App^s

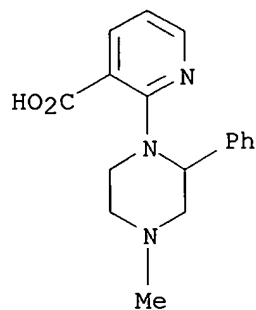
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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	W:				
	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW:				
	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	US 2002165238	A1	20021107	US 2002-73960	20020214
PRAI	US 2001-272699P	P	20010301		
OS	CASREACT 137:216962				
AB	The prepn. of 1-(3-carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine dihydrate (I) and other mirtazapine intermediates are described. These compds. are particularly useful in the prepn. of mirtazapine. Thus, 1-(3-cyano-2-pyridyl)-4-methyl-2-phenylpiperazine was hydrolyzed with aq. KOH, neutralized with HCl and the ppt. washed with water to give I whose crystal structure is reported.				
IT	457601-25-1P , 1-(3-Carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine dihydrate RL: IMF (Industrial manufacture); PRP (Properties); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (prepn. of 1-(3-carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine dihydrate as an intermediate for mirtazapine)				
RN	457601-25-1 CAPLUS				
CN	3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)-, dihydrate (9CI) (CA INDEX NAME)				



2 H₂O

10/073960

IT **61338-13-4P**, 1-(3-Carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine
RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic
preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. of 1-(3-carboxy-2-pyridyl)-4-methyl-2-phenylpiperazine
dihydrate as an intermediate for mirtazapine)
RN 61338-13-4 CAPLUS
CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA
INDEX NAME)



RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/073960

L5 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2002 ACS

AN 2002:368461 CAPLUS

DN 136:369741

TI A novel method for preparation of piperazine and its derivatives

IN Sebastian, Sonny; Patel, Hetal Virendra; Thennati, Rajamannar

PA Sun Pharmaceutical Industries Ltd., India

SO PCT Int. Appl., 23 pp.

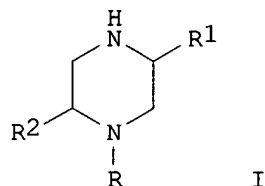
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2002038552	A1	20020516	WO 2001-IN129	20010629
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	AU 2001078669	A5	20020521	AU 2001-78669	20010629
	BE 1013317	A6	20011106	BE 2001-513	20010727
	CH 692342	A	20020515	CH 2001-1428	20010802
	US 2002095038	A1	20020718	US 2001-37309	20011025
PRAI	IN 2000-MU994	A	20001107		
	WO 2001-IN129	W	20010629		
OS	CASREACT 136:369741; MARPAT 136:369741				
GI					



AB Compds. I [R = H, C1-6 alkyl, phenyl-C1-4 alkyl; R1 = H, Me, (un)substituted phenyl; R2 = H, Me, fluoromethyl] useful as starting materials for prepn. of pharmaceutically active compds. are prepd. by reacting R1COCO2R with H2NCH2CHR2NHR to give 3,4-dehydropiperazine-2-one and its derivs., followed by reacting with a reducing agent to yield I. Thus, 1-methyl-3-phenylpiperazine was prepd. and used as starting material for prepn. of 1,2,3,4,10,14b-hexahydro-2-methyl-pyrazino[2,1-a]pyrido[2,3-c][2]benzazepine.

IT **61338-13-4P**

RL: IMF (Industrial manufacture); RCT (Reactant); PREP (Preparation); RACT (Reactant or reagent)

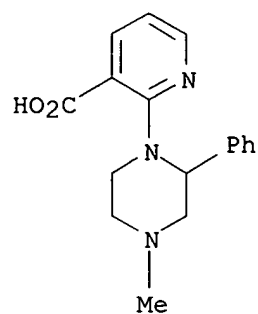
(intermediate; prepn. of piperazine derivs. as starting materials for prepn. of pharmaceutically active compds.)

RN 61338-13-4 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA

10/073960

INDEX NAME)



RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/073960

L5 ANSWER 3 OF 7 CAPLUS COPYRIGHT 2002 ACS

AN 2001:435071 CAPLUS

DN 135:33494

TI Process for the preparation of a pyridinemethanol compound

IN Iishi, Eiichi; Yoshikawa, Kanami

PA Sumika Fine Chemicals Co., Ltd., Japan

SO PCT Int. Appl., 30 pp.

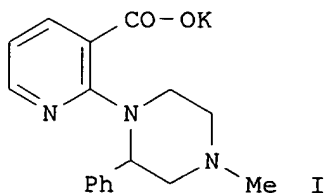
CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 2

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PI	WO 2001042240	A1	20010614	WO 2000-JP6688	20000928
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	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	WO 2001042239	A1	20010614	WO 2000-JP5384	20000811
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	AU 2000074472	A5	20010618	AU 2000-74472	20000928
	EP 1238977	A1	20020911	EP 2000-962909	20000928
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
PRAI	JP 1999-353514	A	19991213		
	WO 2000-JP5384	W	20000811		
	WO 2000-JP6688	W	20000928		
OS	CASREACT 135:33494				
GI					



AB A pyridinemethanol compd. useful as an important intermediate for the prepn. of mirtazapine effective as an antidepressant can be prepd. by ~~reducing a potassium salt of pyridinecarboxylic acid as represented by formula I with a metal hydride.~~ Thus, 1-butanol 162, KOH 60.93, and

10/073960

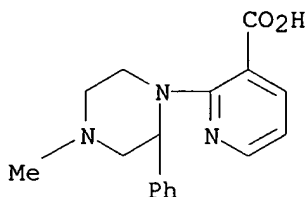
2-(4-methyl-2-phenylpiperazin-1-yl)pyridine-3-carbonitrile oxalate 40 g were heated to give potassium 2-(4-methyl-2-phenylpiperazin-1-yl)pyridine-3-carboxylate, which was reduced in THF with 12.5 g lithium aluminum hydride to give 21.78 g 2-(4-methyl-2-phenylpiperazin-1-yl)pyridine-3-methanol (yield 70.78%).

IT **343626-55-1P**

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; prepn. of pyridinemethanol compd. as intermediate for mirtazapine)

RN 343626-55-1 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)-, potassium salt (9CI) (CA INDEX NAME)



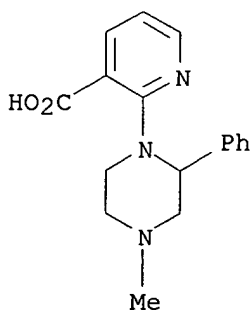
● K

IT **61338-13-4P**

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
(prepn. of pyridinemethanol compd. as intermediate for mirtazapine)

RN 61338-13-4 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA INDEX NAME)



RE.CNT 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/073960

L5 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2002 ACS

AN 2001:435070 CAPLUS

DN 135:33493

TI Process for the preparation of a pyridinemethanol compound

IN Iishi, Eiichi; Yoshikawa, Kanami

PA Sumika Fine Chemicals Co., Ltd., Japan

SO PCT Int. Appl., 30 pp.

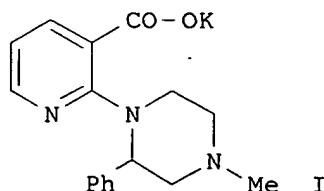
CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 2

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001042239	A1	20010614	WO 2000-JP5384	20000811
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	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	AU 2000064742	A5	20010618	AU 2000-64742	20000811
	WO 2001042240	A1	20010614	WO 2000-JP6688	20000928
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	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	AU 2000074472	A5	20010618	AU 2000-74472	20000928
	EP 1238977	A1	20020911	EP 2000-962909	20000928
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL			
	US 6376668	B1	20020423	US 2000-706803	20001107
	US 2002035255	A1	20020321	US 2001-981919	20011019
	US 6437120	B1	20020820		
PRAI	JP 1999-353514	A	19991213		
	WO 2000-JP5384	W	20000811		
	WO 2000-JP6688	W	20000928		
	US 2000-706803	A3	20001107		
OS	CASREACT 135:33493				
GI					



10/073960

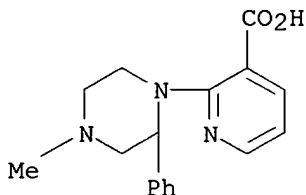
AB A pyridinemethanol compd. serving as an important intermediate of mirtazapine useful as antidepressant can be prepd. by reducing a potassium salt of a pyridinecarboxylic acid as represented by formula I with a metal hydride.

IT **343626-55-1P**

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; prepn. of pyridinemethanol compd. as intermediate for mirtazapine)

RN 343626-55-1 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)-, potassium salt (9CI) (CA INDEX NAME)



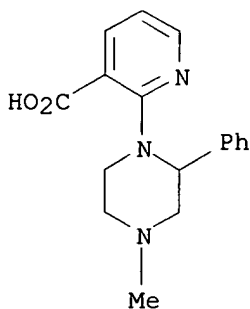
● K

IT **61338-13-4P**

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
(prepn. of pyridinemethanol compd. as intermediate for mirtazapine)

RN 61338-13-4 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA INDEX NAME)



RE.CNT 15 THERE ARE 15 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/073960

L5 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2002 ACS

AN 2000:756528 CAPLUS

DN 133:321900

TI Novel synthesis and crystallization of piperazine ring-containing compounds such as mirtazapine

IN Singer, Claude; Liberman, Anita; Finkelstein, Nina

PA Teva Pharmaceutical Industries Ltd., Israel; Teva Pharmaceuticals Usa, Inc.

SO PCT Int. Appl., 22 pp.

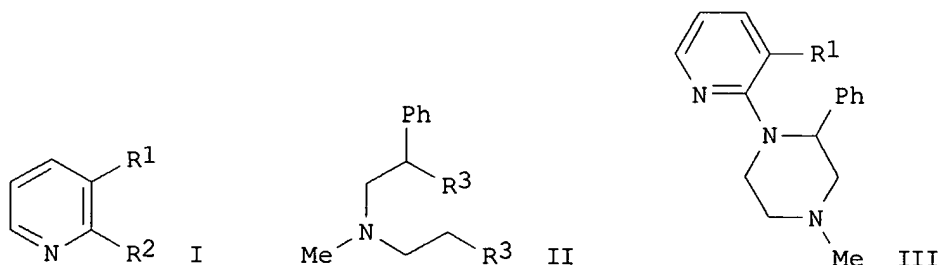
CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2000062782	A1	20001026	WO 2000-US10357	20000418
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	RW:	GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG			
	EP 1178805	A1	20020213	EP 2000-923457	20000418
	R:	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO			
	US 2001051718	A1	20011213	US 2001-900646	20010706
PRAI	US 1999-130047P	P	19990419		
	US 2000-552485	A3	20000418		
	WO 2000-US10357	W	20000418		
OS	CASREACT 133:321900; MARPAT 133:321900				
GI					



AB Mirtazapine, useful in treating depression (no data), was prep'd. by reacting pyridine I [R1 = CH2OH, CH2Cl, CH2Br, CH2I; R2 = NH2] with comp'd. II [R3 = Cl, F, Br, I] followed by treating the resulting piperazine III with H2SO4. The mirtazapine intermediate 1-(3-carboxypyridyl-2)-4-methyl-2-phenylpiperazine may be made by hydrolyzing 1-(3-cyanopyridyl-2)-4-methyl-2-phenylpiperazine with KOH at a temp. of at least about 130.degree.C. The present invention also relates to new processes for recrystn. of mirtazapine from crude mirtazapine.

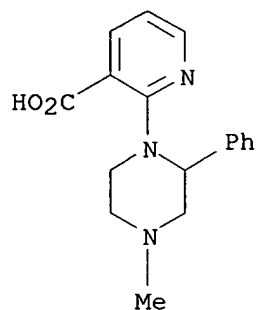
10/073960

IT **61338-13-4P**

RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(novel synthesis and crystn. of piperazine ring-contg. compds. such as mirtazapine)

RN 61338-13-4 CAPLUS

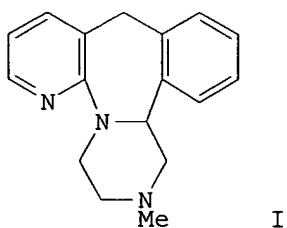
CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA INDEX NAME)



RE.CNT 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

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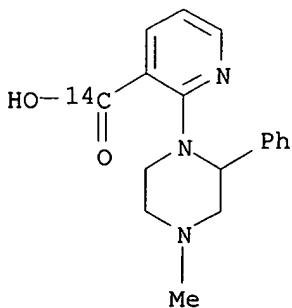
L5 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2002 ACS
AN 1990:139001 CAPLUS
DN 112:139001
TI The synthesis of Org 3770 labeled with tritium, carbon-13 and carbon-14
AU Kaspersen, Frans M.; Van Rooij, Fons A. M.; Sperling, Eric G. M.;
Wieringa, Joop H.
CS Sci. Dev. Group, Organon Int. BV, Oss, 5340 BH, Neth.
SO Journal of Labelled Compounds and Radiopharmaceuticals (1989), 27(9),
1055-68
CODEN: JLCRD4; ISSN: 0362-4803
DT Journal
LA English
OS CASREACT 112:139001
GI



AB The syntheses of 1,2,3,4,10,14b-hexahydro-2-methylpyrazino[2,1-a]pyrido[2,3-c][2]benzazepine (Org 3770, I) labeled with 3H (and 2H), 13C and 14C are described. Tritiated I was prepd. either by exchange under alk. conditions with tritiated water or catalytic reductive dehalogenation of a chloro analog with 3H2. 13C-labeled material was obtained in a 7-step synthesis starting from 13C-labeled benzene, whereas I-14C was prepd. in a 3-step synthesis starting with 14CO2.

IT **125967-25-1P**
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. and redn. of)

RN 125967-25-1 CAPLUS
CN 3-Pyridinecarboxylic-14C acid, 2-(4-methyl-2-phenyl-1-piperaziny)- (9CI)
(CA INDEX NAME)

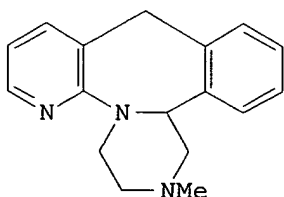


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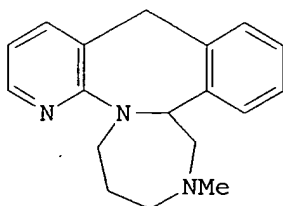
L5 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2002 ACS
AN 1977:29883 CAPLUS
DN 86:29883
TI Heterocyclic tetracyclic compounds
IN Van der Burg, Willem J.
PA AKZO N. V., Neth.
SO Ger. Offen., 40 pp.
CODEN: GWXXBX
DT Patent
LA German
FAN.CNT 1

provided US

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	DE 2614406	A1	19761014	DE 1976-2614406	19760402
	DE 2614406	C2	19920220		
	NL 7504075	A	19761007	NL 1975-4075	19750405
	NL 189199	B	19920901		
	NL 189199	C	19930201		
	ZA 7601756	A	19770330	ZA 1976-1756	19760323
	AU 7612361	A1	19770929	AU 1976-12361	19760325
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	DK 142498	C	19810706		
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	BE 840362	A1	19761004	BE 1976-165832	19760402
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	SE 422941	C	19820715		
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	JP 59042678	B4	19841016		
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	HU 21859	O	19820227	HU 1976-AO437	19760405
	HU 179401	B	19821028		
PRAI	NL 1975-4075		19750405		
GI	NL 1975-75040		19750405		



I



II

AB The title compds., e.g. I and II, with nervous system-depressant and antihistaminic activities (no data), are prepd. by various procedures. Thus, reaction of 2-chloronicotinonitrile with 1-methyl-3-phenylpiperazine gives 2-(4-methyl-2-phenyl-1-piperazinyl)-3-pyridinecarbonitrile which is

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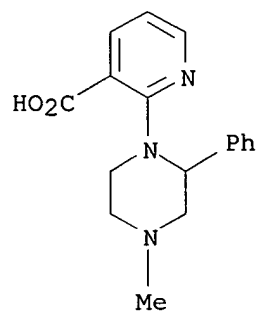
hydrolyzed to the carboxylic acid which is reduced to the hydroxymethyl deriv. (III). Cyclization of 3.25 g III in concd. H₂SO₄ at 20-35.degree. gives after 2 hr and treatment with NH₄OH 2.43 g I.

IT **61338-13-4P**

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(prepn. and redn. of)

RN 61338-13-4 CAPLUS

CN 3-Pyridinecarboxylic acid, 2-(4-methyl-2-phenyl-1-piperazinyl)- (9CI) (CA INDEX NAME)



10/073960

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FULL ESTIMATED COST	33.50	181.83

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
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COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.38	182.21

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